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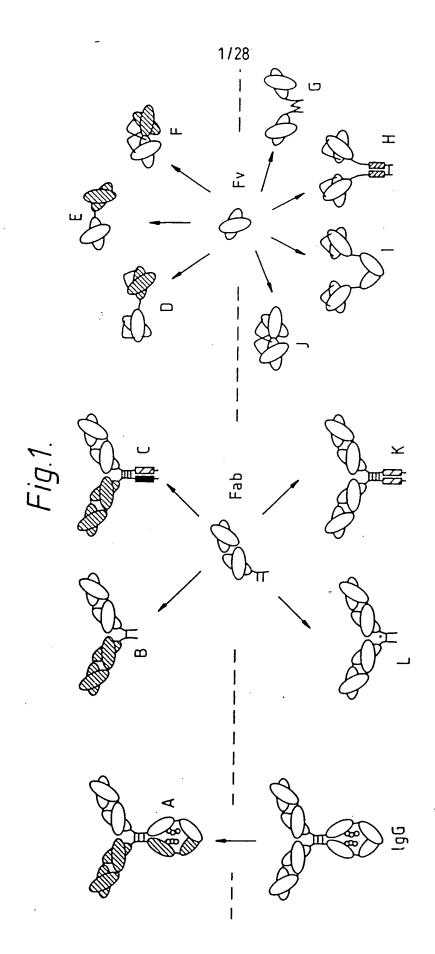
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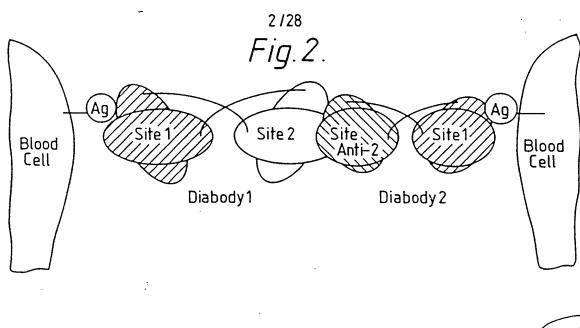
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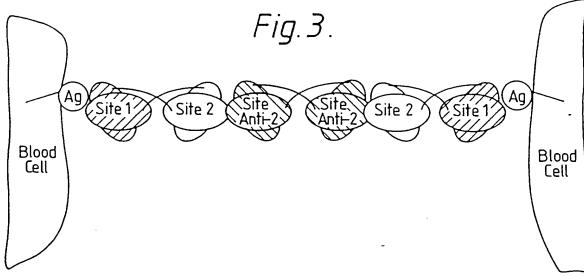
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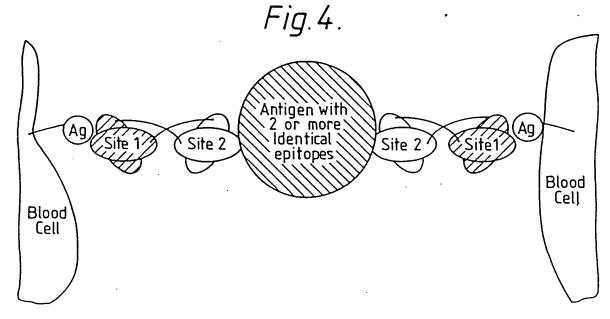


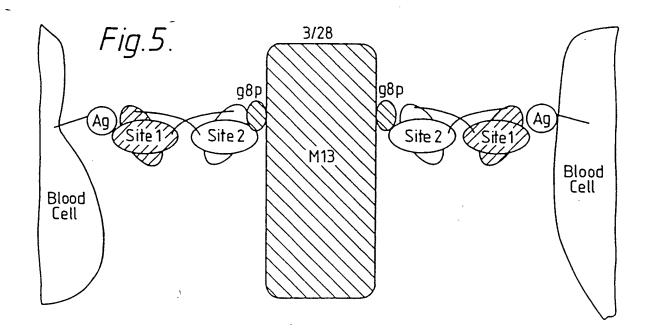
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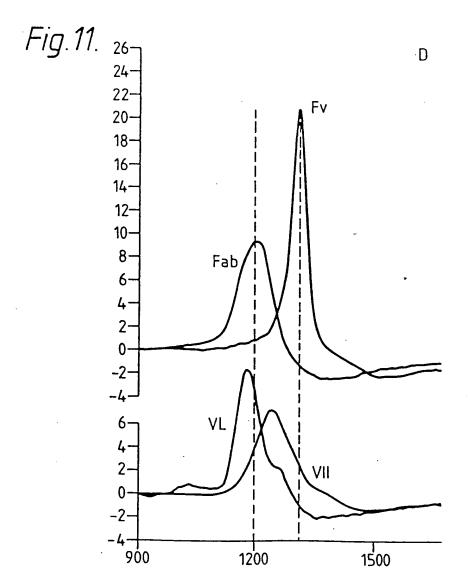
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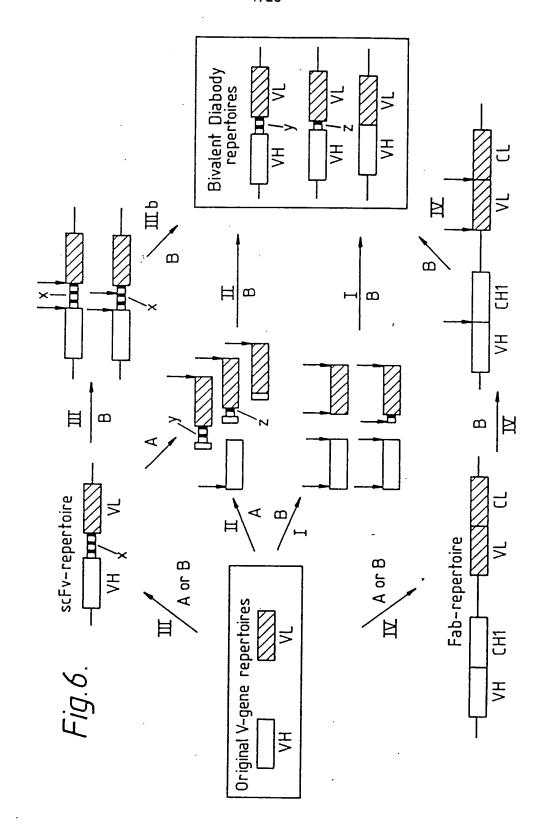








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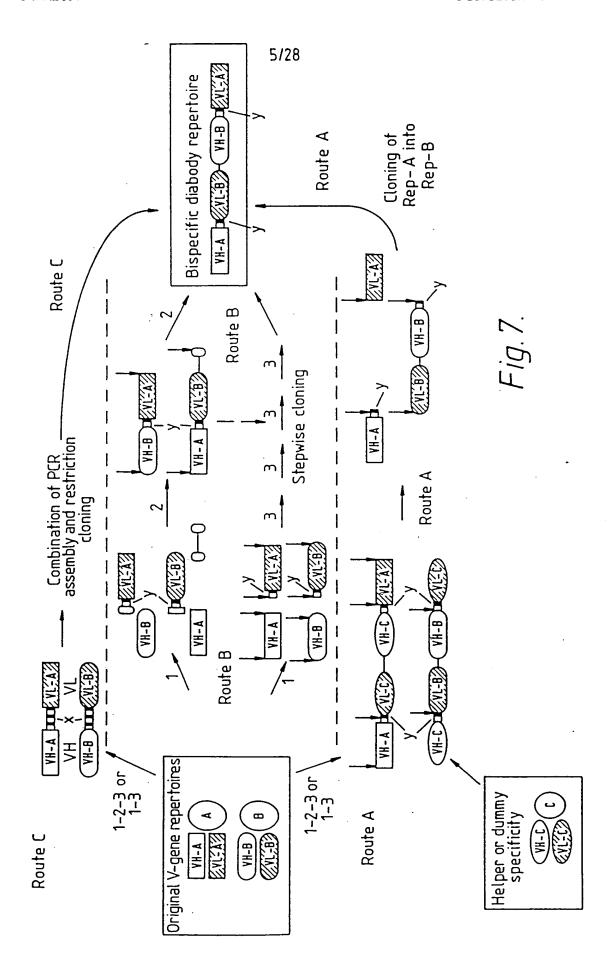
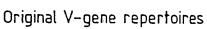
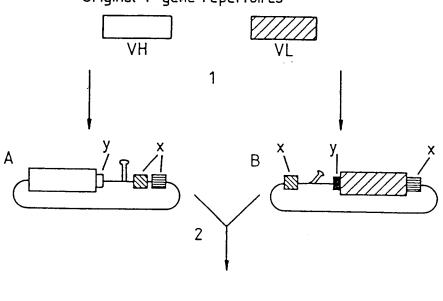
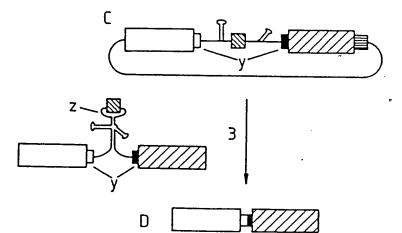
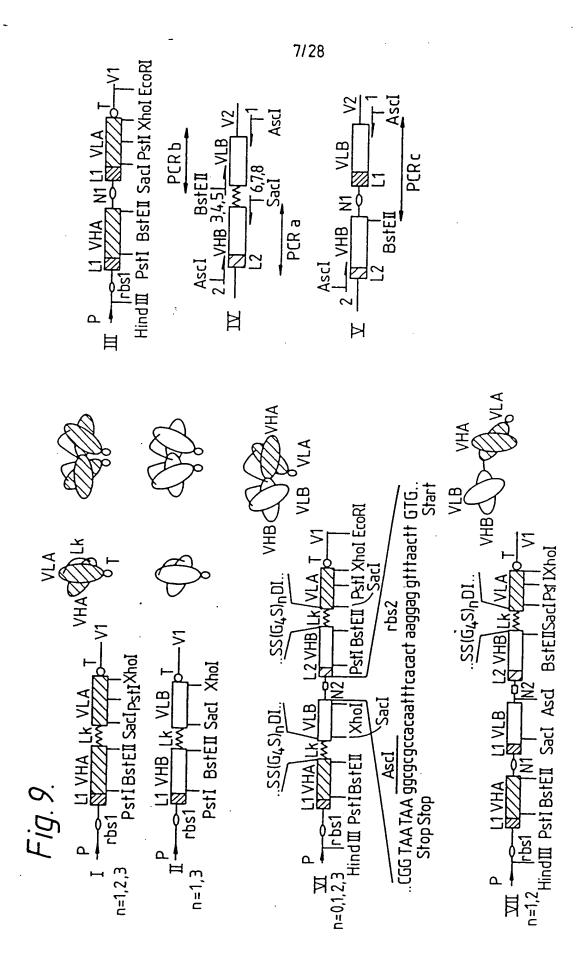


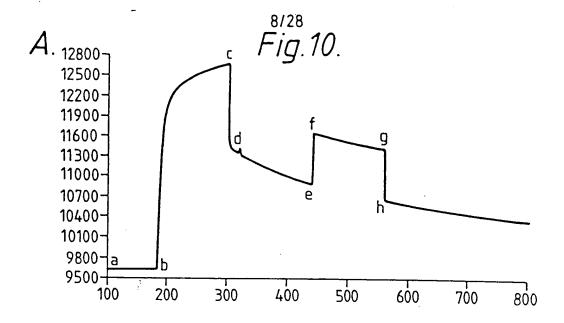
Fig.8.

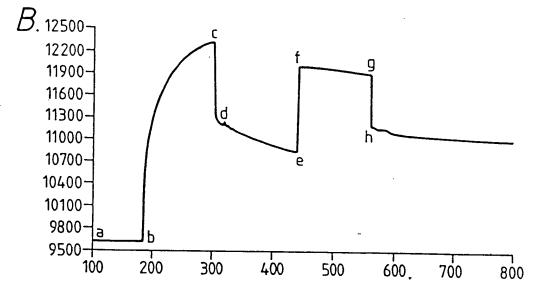












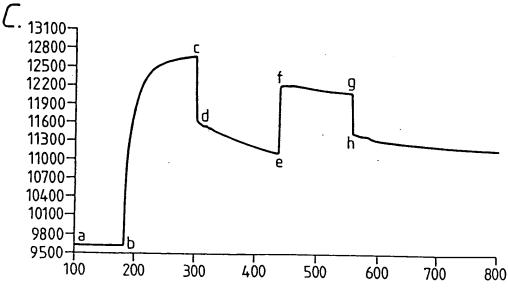
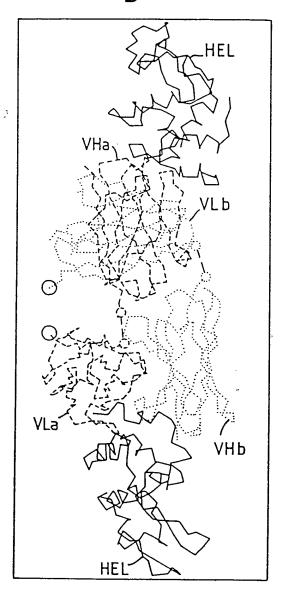
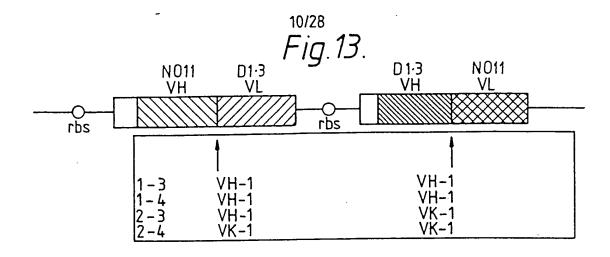
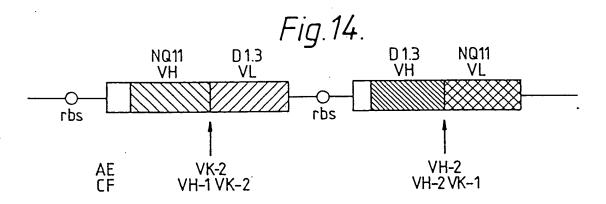


Fig.12.







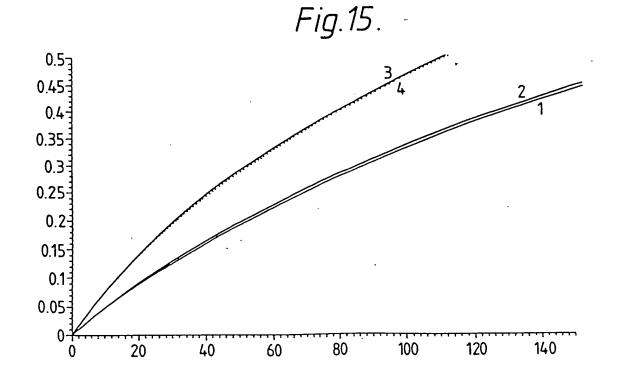
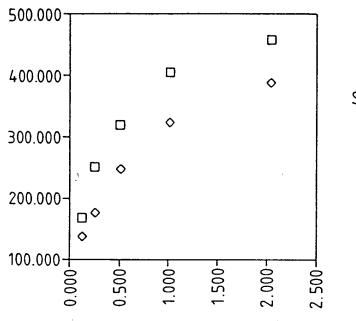


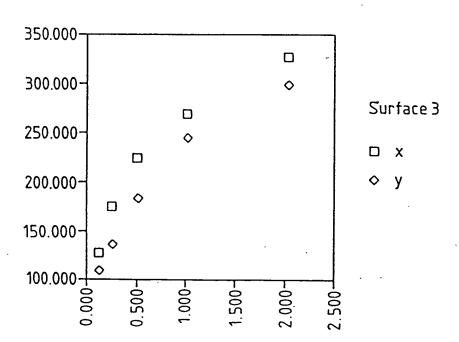
Fig.16.

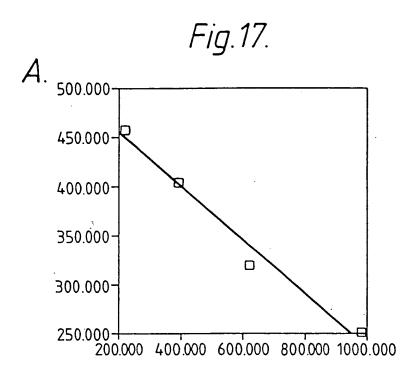


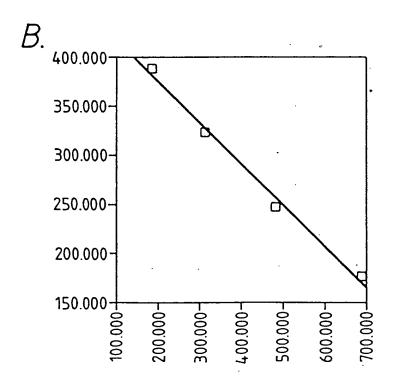
Surface 2

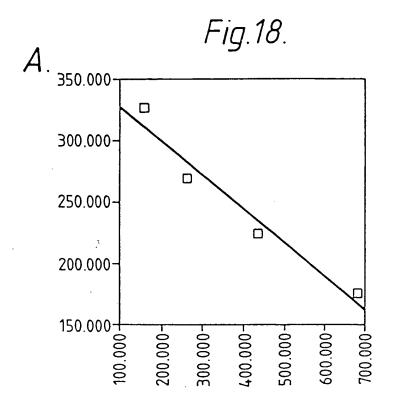
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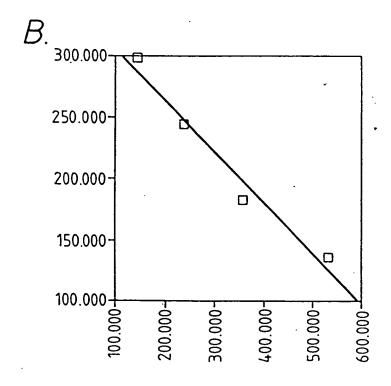
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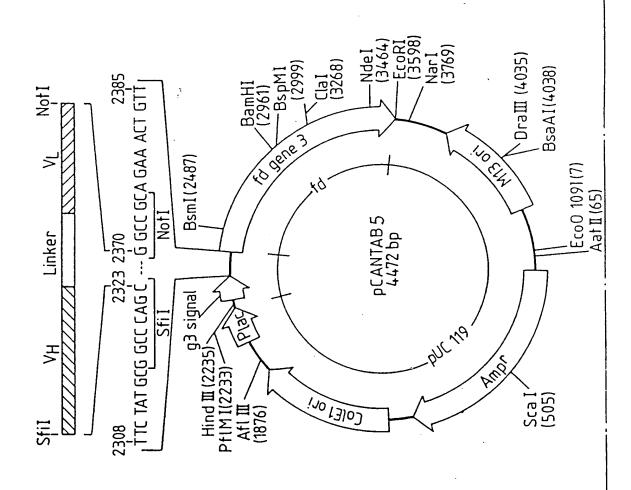


Fig. 19

V K K L L F A I P L V V F Y A A Q P A M A Q V Q L Q V V K K L L F A I P L V V F Y A A Q P A M A Q V Q L Q V GIG AAA AAA TTA TTC GCT ATT CCT TTA GTT GTT TTC TAT GCG GCC CAG GCC CAG GTC CAA CTG CAG GTC FS1 I Sal I Sal I D L E I K R A A E Q K L I S E E D L N G A A (E) T V E GAC CTC GAG GAT CTG AAT GGG GCC GCA TAG ACT GTT GAA Xho I gII structural sfi-Not c-myc gII/pel B hybrid g signal Fig. 19 (Cont). plac

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pCANTAB-5 myc

RECTIFIED SHEET (RULE 91)

HindIII <u>AAGCTT</u> TGGAGCCTTT TTTTGGAGA TTTTCAAC GTG AAA AAA TTA TTC GCA V K K L L F A

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GAA AAT TCA TTT ACT AAC GTC TGG E N S F T N V Fig. 21.

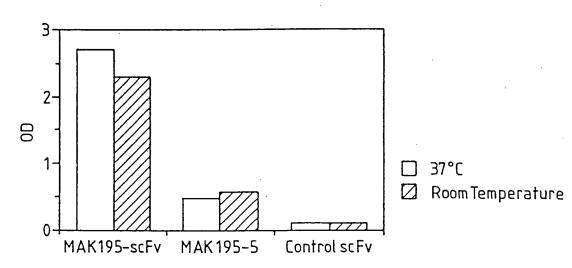
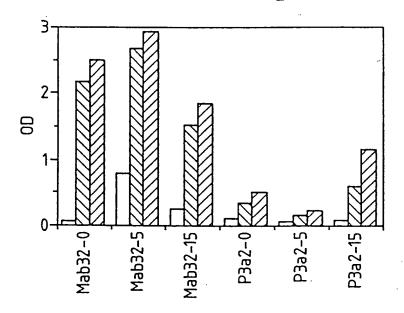


Fig. 22.



- ☐ 37°C
- ZJ 30°,C
- ☑ Room Temp.

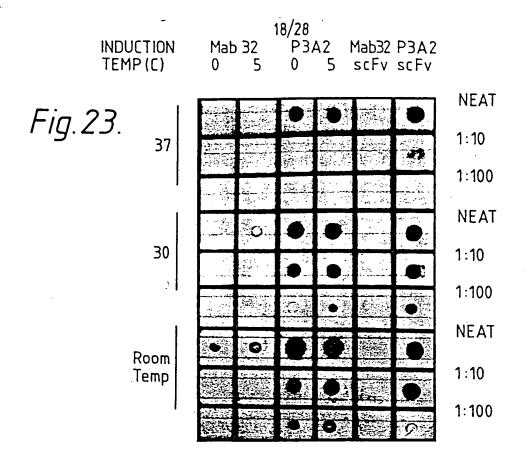
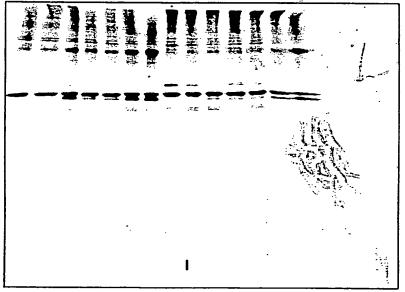


Fig. 24.

1 2 3 4 5 6 7 8 9 10 11 12 13 14



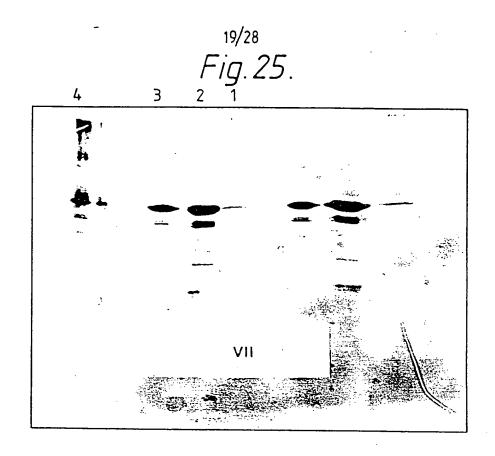
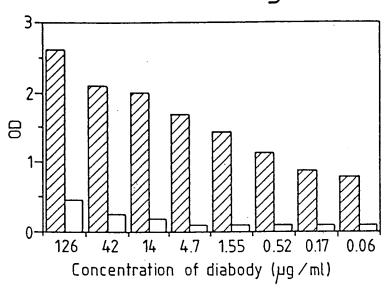


Fig. 26.
0 1 2 3 4 5 6 7 8 9 0.

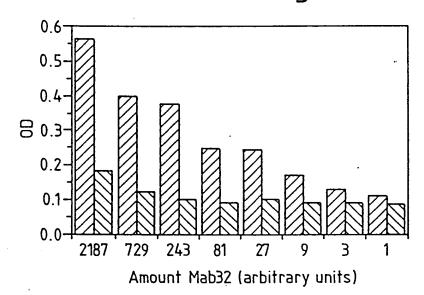
IV

Fig.27.



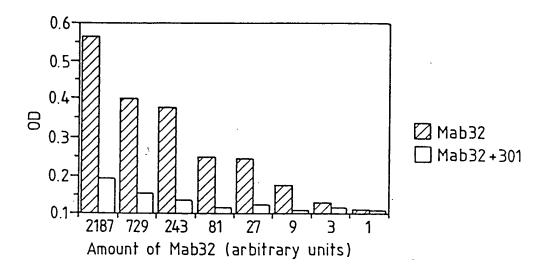
- Diabody on TNF
- ☐ Diabody on BSA

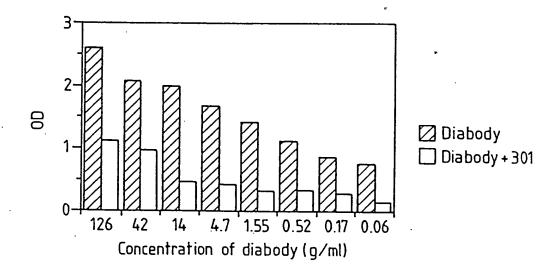
Fig. 28.



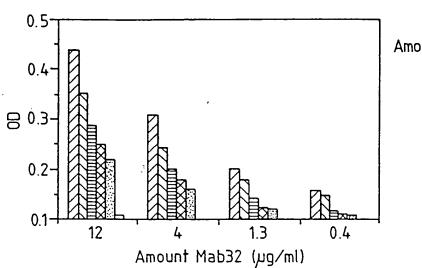
- Mab32
- Mab32+DIABODY

Fig. 29.









## Amount biotin-TNF used (nM)

**250** 

**3** 83.3

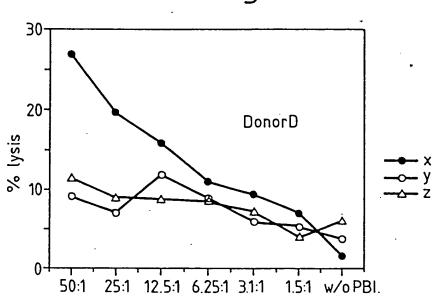
**27.8** 

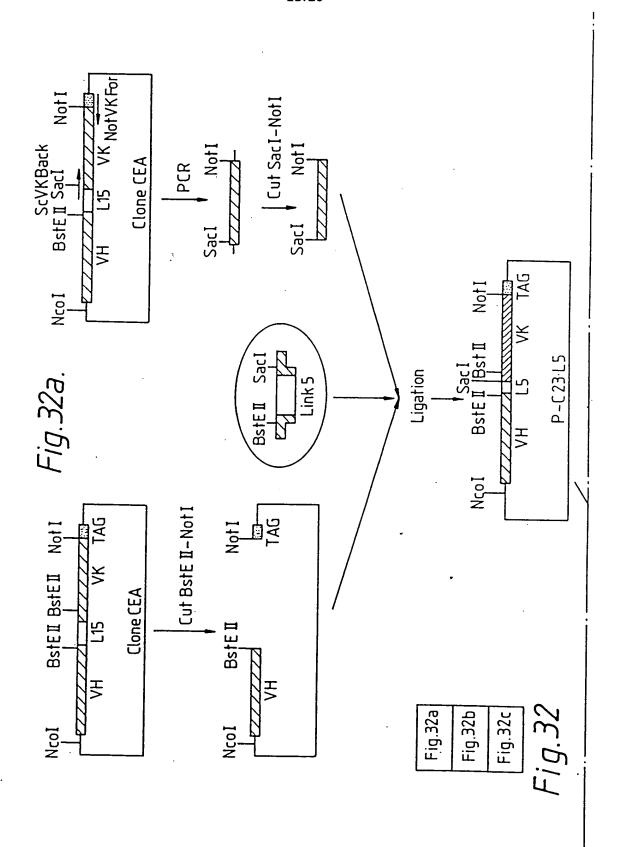
**9.3** 

**3.1** 

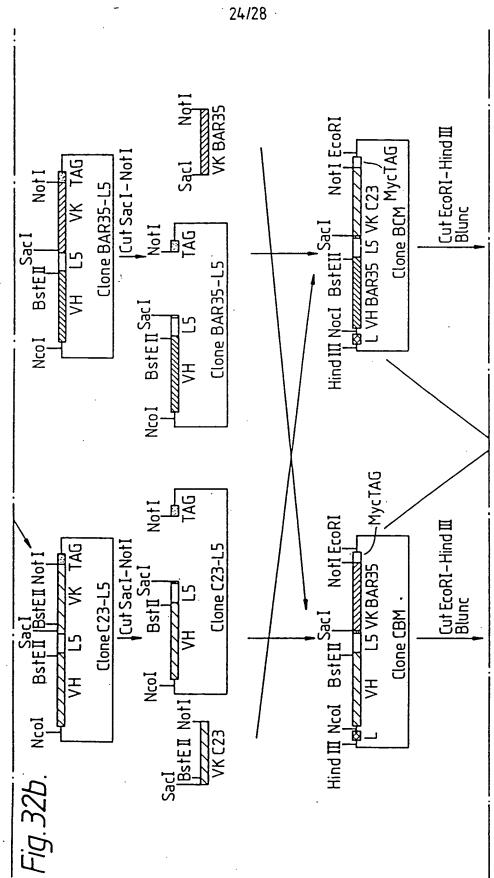
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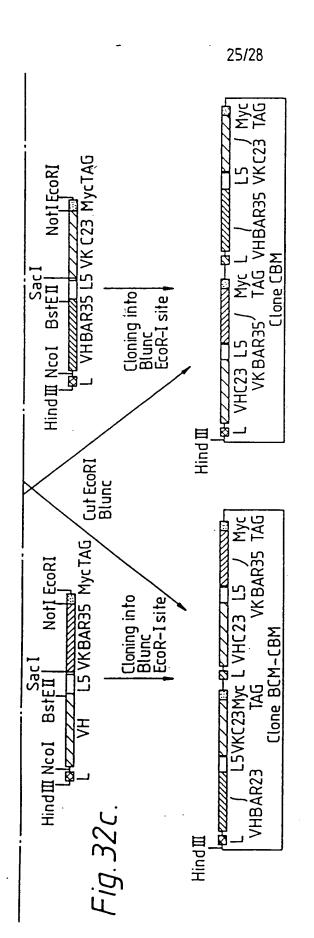
Fig. 31.

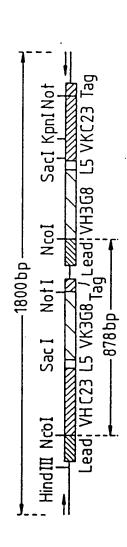






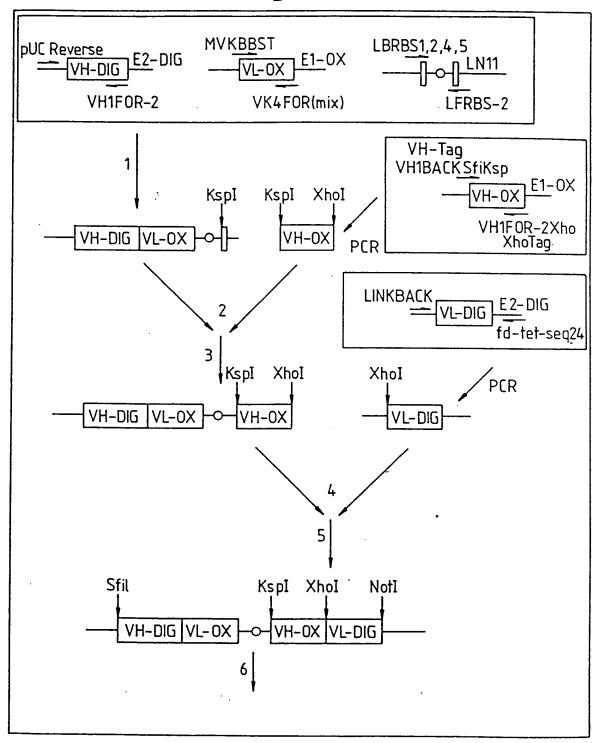


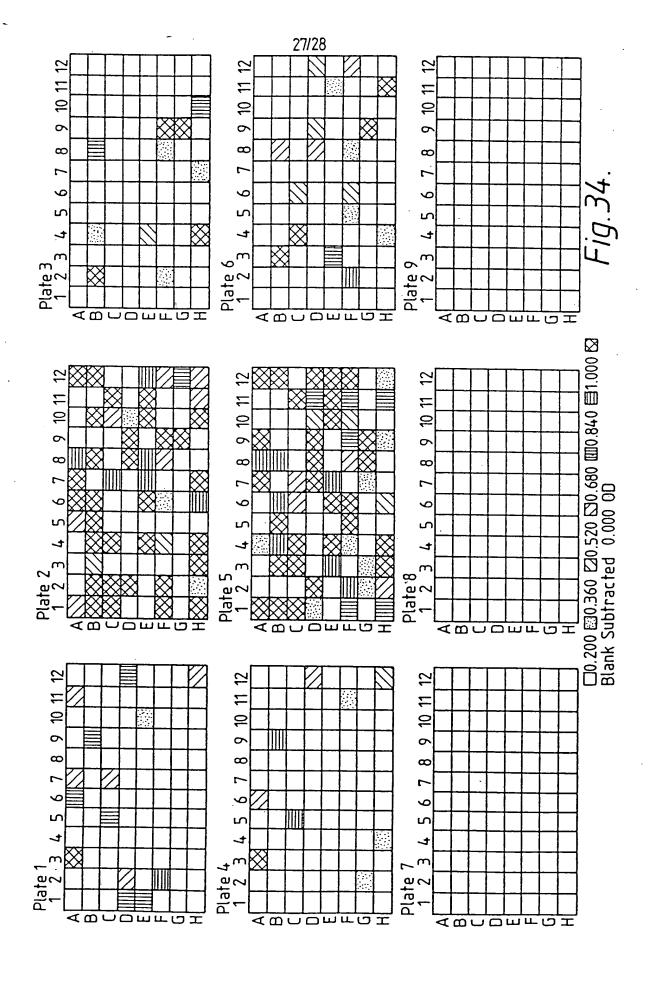


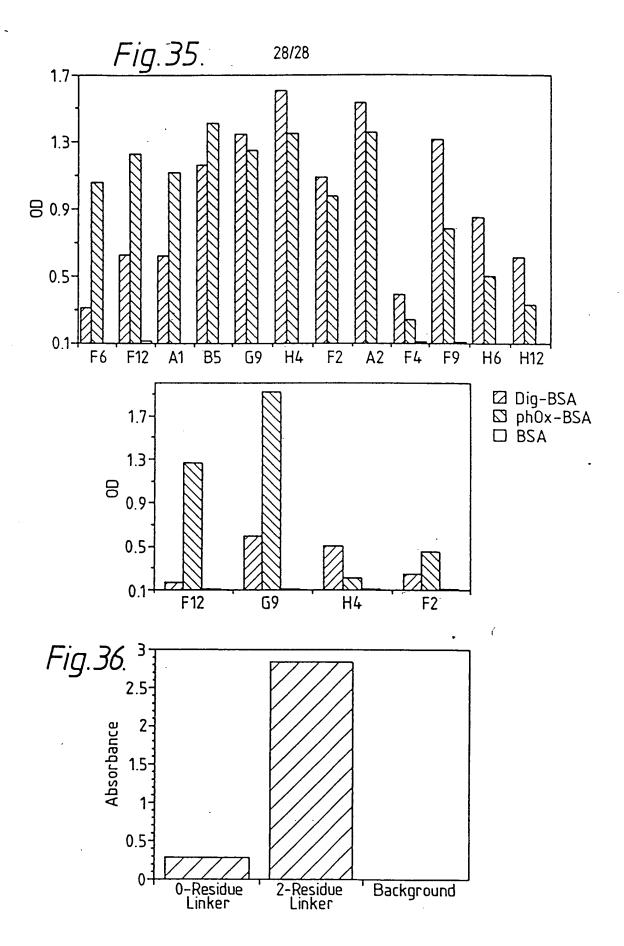


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Fig. 33.







#### INITERNATIONAL SEARCH REPORT

PCT/GB 93/02492

A. CLASSIFICATION OF SUBJECT MATTER
1PC 5 C12N15/13 C07K15/28 C12P21/08 C12N15/62 C12N15/70 C07K1/00 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Mimmur. documentation searched (classification system followed by classification symbols) IPC 5 C12N C07K C12P Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Category \* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. ¥ TRENDS IN BIOTECHNOLOGY 10,11, vol. 10, no. 3, March 1992, CAMBRIDGE, 27,35, 44,45 pages 80 - 84 D. CHISWELL ET AL. 'Phage antibodies: Will new 'colicional' antibodies replace monoclonal antibodies.' see abstract P,X PROCEEDINGS OF THE NATIONAL ACADEMY OF 1-5. SCIENCES OF THE USA 12-37, vol. 90, no. 14 , 15 July 1993 , 39,40, WASHINGTON DC, USA 46-51,60 pages 6444 - 6448 P. HOLLIGER ET AL. '"Diabodies": small bivalent and bispecific antibody fragments.' see the whole document -/--Further documents are listed in the continuation of box C. X Patent family members are listed in annex. Special categories of cited documents: T later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the 'A' document defining the general state of the art which is not considered to be of particular relevance E earlier document but published on or after the international filing date "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such docu-\*O\* document referring to an oral disclosure, use, exhibition or other means ments, such combination being obvious to a person skilled document published prior to the international filing date but later than the priority date claimed '&' document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 20. 05. 94 4 May 1994 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Ripwijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax (+31-70) 340-3016 Nooij, F

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INTERNATIONAL SEARCH RELIVAT

Information on patent family members

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WO-A-9311161 10-06-93 AU-A- 3178993 28-06-93